ristopher Eckman et al. Attorney's Docket No.: 07039-235001

Applicant: Christopher Eckman et al.

Serial No.: 09/824,924 Filed: April 3, 2001

Page :

linked secondary antibody (Amersham, Uppsala, Sweden) using ECL™ Western blotting reagents (Amersham) followed by exposure to X-ray film.--

Please replace the Sequence Listing submitted on November 26, 2001 with the enclosed copy of the Sequence Listing.

## In the Claims:

Please replace claim 7 with amended claim 7 below. For convenience, a full set of pending claims is shown below.

--7. (Twice Amended) A method of identifying a compound that increases the activity of an endothelin converting enzyme (ECE) polypeptide, the method comprising:

contacting  $A\beta$  with an ECE polypeptide in the presence of said compound; and detecting the amount of unhydrolyzed  $A\beta$ ,

wherein a decrease in the amount of unhydrolyzed Aβ produced in the presence of said compound compared to the amount of unhydrolyzed Aβ produced in the absence of said compound is an indication that said compound increases the activity of an ECE polypeptide.--

- 8. The method of claim 7, wherein said ECE and said  $A\beta$  are in a cell.
- 9. The method of claim 7, wherein said unhydrolyzed  $A\beta$  is detected using an immunoassay.
- 11. A method of identifying a compound that has anti-hypertension activity but does not cause an increase in the level of  $A\beta$ , the method comprising:

contacting  $A\beta$  with an ECE in the presence of said compound;

detecting the amount of unhydrolyzed  $A\beta$ , wherein lack of an increase in the amount of unhydrolyzed  $A\beta$  produced in the presence of said compound compared to the amount of unhydrolyzed  $A\beta$  produced in the absence of said compound is an indication that said compound does not cause an increase in the level of said ECE; and

determining the anti-hypertension activity of said compound.

12. The method of claim 11, wherein the anti-hypertension activity of said compound is determined in an animal.

05

Applicant: Christopher Eckman et al. Attorney's Docket No.: 07039-235001

Serial No. : 09/824,924 Filed : April 3, 2001

Page : 5

13. The method of claim 12, wherein said animal is a spontaneously hypersensitive rat (SHR).

14. A method of determining that an anti-hypertension compound or candidate compound does not cause an increase in the level of  $A\beta$ , the method comprising:

contacting  $A\beta$  with an ECE in the presence of said anti-hypertension compound or candidate compound; and

detecting the amount of unhydrolyzed AB,

wherein the lack of an increase in the amount of unhydrolyzed  $A\beta$  produced in the presence of said compound compared to the amount of unhydrolyzed  $A\beta$  produced in the absence of said compound is an indication that said compound does not cause an increase in the level of said ECE.

- 15. The method of claim 14, wherein said anti-hypertension compound or candidate compound is an ECE inhibitor.
- 40. The method of claim 8, wherein said cell is selected from the group consisting of H4 neuroglioma cells, CHO cells, and HUVEC cells.
- 41. The method of claim 7, wherein said compound is selected from the group consisting of a nucleic acid, a polypeptide, a chemical compound, a bacterial extract, a fungal extract, and a plant extract.
- 42. The method of claim 12, wherein said unhydrolyzed  $A\beta$  is detected in said animal.
- 43. The method of claim 11, wherein said unhydrolyzed  $A\beta$  is detected using an immunoassay.
- 44. The method of claim 11, wherein said compound is selected from the group consisting of a nucleic acid, a polypeptide, a chemical compound, a bacterial extract, a fungal extract, and a plant extract.
- 45. The method of claim 14, wherein said unhydrolyzed  $A\beta$  is detected using an immunoassay.
  - 46. The method of claim 14, wherein said unhydrolyzed  $A\beta$  is detected in an animal.
  - 47. The method of claim 46, wherein said animal is a SHR.

Applicant: Christopher Eckman et al.

Serial No.: 09/824,924 : Aprii 3, 2001 Filed

Page

48. The method of claim 14, wherein said compound is selected from the group consisting of a nucleic acid, a polypeptide, a chemical compound, a bacterial extract, a fungal extract, and a plant extract.

Attorney's Docket No.: 07039-235001